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NEWS 1 Web Page for STN Seminar Schedule - N. America
 NEWS 2 NOV 21 CAS patent coverage to include exemplified prophetic
 substances identified in English-, French-, German-,
 and Japanese-language basic patents from 2004-present
 NEWS 3 NOV 26 MARPAT enhanced with FSORT command
 NEWS 4 NOV 26 CHEMSAFE now available on STN Easy
 NEWS 5 NOV 26 Two new SET commands increase convenience of STN
 searching
 NEWS 6 DEC 01 ChemPort single article sales feature unavailable
 NEWS 7 DEC 12 GBFULL now offers single source for full-text
 coverage of complete UK patent families
 NEWS 8 DEC 17 Fifty-one pharmaceutical ingredients added to PS
 NEWS 9 JAN 06 The retention policy for unread STNmail messages
 will change in 2009 for STN-Columbus and STN-Tokyo
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 Classification Data
 NEWS 11 FEB 02 Simultaneous left and right truncation (SLART) added
 for CERAB, COMPUAB, ELCOM, and SOLIDSTATE
 NEWS 12 FEB 02 GENBANK enhanced with SET PLURALS and SET SPELLING
 NEWS 13 FEB 06 Patent sequence location (PSL) data added to USGENE
 NEWS 14 FEB 10 COMPENDEX reloaded and enhanced
 NEWS 15 FEB 11 WTEXTILES reloaded and enhanced
 NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3,
 AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.
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* * * * * STN Columbus * * * * *

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=> file caplus

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	ENTRY	SESSION
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FILE 'CAPLUS' ENTERED AT 12:50:23 ON 12 FEB 2009

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FILE COVERS 1907 - 12 Feb 2009 VOL 150 ISS 7
FILE LAST UPDATED: 11 Feb 2009 (20090211/ED)

Caplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2008.

CAS Information Use Policies apply and are available at:

<http://www.cas.org/legal/infopolicy.html>

This file contains CAS Registry Numbers for easy and accurate substance identification.

```
=> antibody
    342815 ANTIBODY
    413274 ANTIBODIES
L1    546015 ANTIBODY
      (ANTIBODY OR ANTIBODIES)

=> monoclonal
    159505 MONOCLONAL
    560 MONOCLONALS
L2    159578 MONOCLONAL
      (MONOCLONAL OR MONOCLONALS)

=> single chain
    1483267 SINGLE
    3557 SINGLES
    1486279 SINGLE
      (SINGLE OR SINGLES)
    802222 CHAIN
    342735 CHAINS
    1004346 CHAIN
      (CHAIN OR CHAINS)
L3    15140 SINGLE CHAIN
      (SINGLE(W)CHAIN)

=> fragment
    222617 FRAGMENT
    215174 FRAGMENTS
L4    381669 FRAGMENT
      (FRAGMENT OR FRAGMENTS)

=> coronavirus
L5    0 CORONIVIRUS

=> coronavirus
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5817 CORONAVIRUS
 1038 CORONAVIRUSES
 L6 5940 CORONAVIRUS
 (CORONAVIRUS OR CORONAVIRUSES)

=> SARS

L7 4917 SARS

=> L1 and L2

L8 155125 L1 AND L2

=> L1

342815 ANTIBODY

413274 ANTIBODIES

L9 546015 ANTIBODY

(ANTIBODY OR ANTIBODIES)

=> L6 and L1

L10 1537 L6 AND L1

=> L2 and L10

L11 495 L2 AND L10

=> L7 and L1

L12 1026 L7 AND L1

=> L12 and L2

L13 238 L12 AND L2

=> nucleoprotein

8472 NUCLEOPROTEIN

6679 NUCLEOPROTEINS

L14 11936 NUCLEOPROTEIN

(NUCLEOPROTEIN OR NUCLEOPROTEINS)

=> L14 and L11

L15 15 L14 AND L11

=> L14 and L13

L16 8 L14 AND L13

=> L5 and L14

L17 0 L5 AND L14

=> L6 and L14

L18 98 L6 AND L14

=> L7 and L14

L19 47 L7 AND L14

=> L2 and L18

L20 16 L2 AND L18

=> L2 and L19

L21 8 L2 AND L19

=> D L21 IEIS ABS 1-8

L21 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2007:147094 CAPLUS
 DOCUMENT NUMBER: 147:46687
 TITLE: Cloning, expressing and antigenicity analysis of nucleocapsid proteins of SARS-CoV, HCoV-229E and OC43
 AUTHOR(S): Che, Xiaoyan; Liao, Zhiyong; Wang, Yadi; Qiu, Liwen; Wen, Kun; Pan, Yuxian; Xu, Hua; Mei, Yabo; Hao, Wei; Ding, Yanqing
 CORPORATE SOURCE: Zhujiang Hospital, Southern Medical University, Guangzhou, 510282, Peop. Rep. China
 SOURCE: Zhonghua Weishengwuxue He Mianyixue Zazhi (2005), 25(9), 711-715
 CODEN: ZWMZDP; ISSN: 0254-5101
 PUBLISHER: Beijing Shengwu Zhipin Yanjiusuo
 DOCUMENT TYPE: Journal
 LANGUAGE: Chinese

AB A recombinant nucleocapsid (N) protein of SARS-CoV, HCoV-229E and HCoV-OC43, was obtained resp. to study antigenic relationships of N proteins between SARS-CoV and human coronaviruses 229E and OC43. The genes encoding the full-length of N proteins from SARS-CoV, HCoV-229E and HCoV-OC43 were amplified by RT-PCR and cloned into the prokaryotic expression vector pQE30. The His6-tagged N proteins were expressed in the M15 strain and further purified with affinity chromatog. The antigenicity of N proteins was analyzed by Western blot and immunofluorescence assay. The N genes of 1281, 1182 and 1359 bp from SARS-CoV, HCoV-229E and HCoV-OC43, resp. were amplified with their corresponding primer pairs. The recombinant plasmids were sequenced, and they were all in frame with sequences matching those for the N genes of the three coronaviruses. The expressed recombinant His6-tagged N proteins were identified by Western blot assay with anti-His tag monoclonal antibody. The immunoreactive protein bands with expected sizes were 47 kDa, 44 kDa and 50 kDa from SARS-CoV, HCoV-229E and HCoV-OC43, resp. The nucleocapsid proteins of SARS-CoV, HCoV-229E and HCoV-OC43 strongly and specifically reacted with the virus specific rabbit serum and with the nucleoprotein specific murine serum. No cross-reactivity was found among the nucleocapsid proteins of SARS-CoV, HCoV-229E and HCoV-OC43. The immunogenic nucleocapsid recombinant proteins from SARS-CoV, HCoV-229E and HCoV-OC43 were obtained. There was no antigenic relationship among the three N proteins.

L21 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:1263772 CAPLUS
 DOCUMENT NUMBER: 146:137159
 TITLE: Coronavirus nucleocapsid protein is an RNA chaperone
 AUTHOR(S): Zuniga, Sonia; Sola, Isabel; Moreno, Jose L.; Sabella, Patricia; Plana-Duran, Juan; Enjuanes, Luis
 CORPORATE SOURCE: Centro Nacional de Biotecnologia, CSIC, Department of Molecular and Cell Biology, Campus Universidad Autonoma, Madrid, 28049, Spain
 SOURCE: Virology (2007), 357(2), 215-227
 CODEN: VIRLAX; ISSN: 0042-6822
 PUBLISHER: Elsevier
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB RNA chaperones are nonspecific nucleic acid binding proteins with long disordered regions that help RNA mols. to adopt its functional conformation. Coronavirus nucleoproteins (N) are nonspecific

RNA-binding proteins with long disordered regions. Therefore, we investigated whether transmissible gastroenteritis coronavirus (TGEV) N protein was an RNA chaperone. Purified N protein enhanced hammerhead ribozyme self-cleavage and nucleic acids annealing, which are properties that define RNA chaperones. In contrast, another RNA-binding protein, PTB, did not show these activities. N protein chaperone activity was blocked by specific monoclonal antibodies. Therefore, it was concluded that TGEV N protein is an RNA chaperone. In addn., we have shown that purified severe acute respiratory syndrome (SARS)-CoV N protein also has RNA chaperone activity. In silico predictions of disordered domains showed a similar pattern for all coronavirus N proteins evaluated. Altogether, these data led us to suggest that all coronavirus N proteins might be RNA chaperones.

REFERENCE COUNT: 83 THERE ARE 83 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:1139612 CAPLUS
 DOCUMENT NUMBER: 146:140843
 TITLE: Antigenic and cellular localisation analysis of the severe acute respiratory syndrome coronavirus nucleocapsid protein using monoclonal antibodies
 AUTHOR(S): Bussmann, Bianca M.; Reiche, Sven; Jacob, Lotta H.; Braun, Jan Matthias; Jassoy, Christian
 CORPORATE SOURCE: Institute of Virology, University of Leipzig, Leipzig, 04103, Germany
 SOURCE: Virus Research (2006), 122(1-2), 119-126
 CODEN: VIREDF; ISSN: 0168-1702
 PUBLISHER: Elsevier B.V.
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB A member of the family of coronaviruses has previously been identified as the cause of the severe acute respiratory syndrome (SARS). In this study, several monoclonal antibodies against the nucleocapsid protein have been generated to examine distribution of the nucleocapsid in virus-infected cells and to study antigenic regions of the protein. Confocal microscopic anal. identified nucleocapsids packaged in vesicles in the perinuclear area indicating viral synthesis at the endoplasmic reticulum and Golgi app. The monoclonal antibodies bound to the central and carboxyterminal half of the nucleocapsid protein indicating prominent exposure and immunogenicity of this part of the protein. Antibodies recognized both linear and conformational epitopes. Predictions of antigenicity using amt.. modeling based on hydrophobicity anal. of SARS nucleoprotein could not be confirmed fully. Antibody binding to discontinuous peptides provides evidence that amino acids 274-283 and 373-382 assemble to a structural unit particularly rich in basic amino acids. In addn., amino acids 286-295, 316-325 and 361-367 that represent the epitope recognized by monoclonal antibody 6D11C1 converge indicating a well-structured C-terminal region of the SARS virus nucleocapsid protein and functional relation of the peptide regions involved. Alternatively, dimerization of the nucleocapsid protein may result in juxtaposition of the amino acid sequences 316-325 and 361-367 on one nucleoprotein mol. to amino acid 286-295 on the second peptide. The monoclonal antibodies will be available to assess antigenicity and immunol. variabilities between different SARS CoV strains.

REFERENCE COUNT: 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:236188 CAPLUS
 DOCUMENT NUMBER: 145:185561
 TITLE: Time course and cellular localization of SARS-CoV nucleoprotein and RNA in lungs from fatal cases of SARS
 AUTHOR(S): Nicholls, John M.; Butany, Jagdish; Poon, Leo L. M.; Chan, Kwok H.; Beh, Swan Lip; Poutanen, Susan; Peiris, J. S. Malik; Wong, Maria
 CORPORATE SOURCE: Department of Pathology, The University of Hong Kong, Pok Fu Lam, Hong Kong SAR, Peop. Rep. China
 SOURCE: PLoS Medicine (2006), 3(2), 222-229
 CODEN: PMLEAC; ISSN: 1549-1277
 URL: <http://medicine.plosjournals.org/archive/1549-1676/3/2/pdf/10.1371.1549-1676.3.2.complete.pdf>
 PUBLISHER: Public Library of Science
 DOCUMENT TYPE: Journal; (online computer file)
 LANGUAGE: English

AB Background: Cellular localization of severe acute respiratory syndrome coronavirus (SARS-CoV) in the lungs of patients with SARS is important in confirming the etiol. assocn. of the virus with disease as well as in understanding the pathogenesis of the disease. To our knowledge, there have been no comprehensive studies investigating viral infection at the cellular level in humans. Methods and Findings: We collected the largest series of fatal cases of SARS with autopsy material to date by merging the pathol. material from two regions involved in the 2003 worldwide SARS outbreak in Hong Kong, China, and Toronto, Canada. We developed a monoclonal antibody against the SARS-CoV nucleoprotein and used it together with in situ hybridization (ISH) to analyze the autopsy lung tissues of 32 patients with SARS from Hong Kong and Toronto. We compared the results of these assays with the pulmonary pathologies and the clin. course of illness for each patient. SARS-CoV nucleoprotein and RNA were detected by immunohistochem. and ISH, resp., primarily in alveolar pneumocytes and, less frequently, in macrophages. Such localization was detected in four of the seven patients who died within two weeks of illness onset, and in none of the 25 patients who died later than two weeks after symptom onset. Conclusions: The pulmonary alveolar epithelium is the chief target of SARS-CoV, with macrophages infected subsequently. Viral replication appears to be limited to the first two weeks after symptom onset, with little evidence of continued widespread replication after this period. If antiviral therapy is considered for future treatment, it should be focused on this two-week period of acute clin. disease.

REFERENCE COUNT: 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2005:409560 CAPLUS
 DOCUMENT NUMBER: 142:462283
 TITLE: Monoclonal antibodies specific to SARS virus nucleoprotein for immunodiagnosis of SARS
 INVENTOR(S): Uchida, Yoshiaki; Fujii, Nobuyuki; Kurano, Yoshihiro; Okada, Masahisa; Kogaki, Hiroyuki; Kido, Yasuji; Miyake, Kazushige
 PATENT ASSIGNEE(S): Fujirebio Inc., Japan
 SOURCE: PCT Int. Appl., 41 pp.

CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005042579	A1	20050512	WO 2004-JP16099	20041029
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CN 1902230	A	20070124	CN 2004-80039648	20041029
IN 2006KN01457	A	20070504	IN 2006-KN1457	20060530
US 20080254440	A1	20081016	US 2007-577310	20070222
PRIORITY APPLN. INFO.:			JP 2003-373779	A 20031031
			JP 2004-34268	A 20040210
			WO 2004-JP16099	W 20041029

AB Provided are monoclonal antibodies specific to SARS virus nucleoprotein and hybridomas producing them. These monoclonal antibodies are labeled with enzyme and used for immunodiagnosis of SARS.

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:872877 CAPLUS
 DOCUMENT NUMBER: 141:378847
 TITLE: Endogenous host elements or viral-based sequence elements for diagnosis, prognosis and therapy of viral infection, autoimmune disease and lymphoproliferative disease
 INVENTOR(S): Hu, Yu-wen; Brown, Earl
 PATENT ASSIGNEE(S): Canadian Blood Services, Can.
 SOURCE: PCT Int. Appl., 174 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004090544	A2	20041021	WO 2004-CA544	20040413
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,				

BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
 ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
 SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
 TD, TG

CA 2522067 A1 20041021 CA 2004-2522067 20040413
 EP 1625402 A2 20060215 EP 2004-726942 20040413

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR

US 20060115875 A1 20060601 US 2005-248008 20051011

PRIORITY APPLN. INFO.:

US 2003-461137P P 20030409

US 2003-506779P P 20030930

WO 2004-CA544 W 20040413

AB A method of detecting, characterizing and treating viral infection, autoimmune disease and lymphoproliferative disease is provided. In particular, a strategy of mol. mimicry is provided for characterizing viral behavior and/or a predisposition for a given viral outcome in vivo. Novel compns. are also provided for detecting, characterizing and treating viral infections. The viral infection is caused by HCV, HIV, HTLV-1, HTLV-2, SARS-CoV, or a member of Retroviridae, Flaviviridae, Herpesviridae, Papillomaviridae, Poxviridae or Coronaviridae. The viral-based sequence element is e.g. an element of S protein sequence of an ORF1a protein sequence of SARS-CoV; a Gag, Pol or Env polypeptide of HTLV-1; a NS5A and E2 protein of HCV; bacterial virulence factor; human endogenous retrovirus element; Peyer's patches virulence factor gipA; or an Ig selected from IgG, IgA, IgM, IgD or IgE. The treatment regime includes an anti-viral monoclonal or polyclonal antibody, or a compd. capable of binding epitope of the endogenous host element.

L21 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER:

2004:541777 CAPLUS

DOCUMENT NUMBER:

141:222968

TITLE:

Organ distribution of severe acute respiratory syndrome (SARS) associated coronavirus (SARS-CoV) in SARS patients: implications for pathogenesis and virus transmission pathways

AUTHOR(S):

Ding, Yanqing; He, Li; Zhang, Qingling; Huang, Zhongxi; Che, Xiaoyan; Hou, Jinlin; Wang, Huijun; Shen, Hong; Qiu, Liwen; Li, Zhuguo; Geng, Jian; Cai, Junjie; Han, Huixia; Li, Xin; Kang, Wei; Weng, Desheng; Liang, Ping; Jiang, Shibo

CORPORATE SOURCE:

Department of Pathology, Nan Fang Hospital, First Military Medical University, Guangzhou, Peop. Rep. China

SOURCE:

Journal of Pathology (2004), 203(2), 622-630

CODEN: JPTLAS; ISSN: 0022-3417

PUBLISHER:

John Wiley & Sons Ltd.

DOCUMENT TYPE:

Journal

LANGUAGE:

English

AB We previously identified the major pathol. changes in the respiratory and immune systems of patients who died of severe acute respiratory syndrome (SARS) but gained little information on the organ distribution of SARS-assocd. coronavirus (SARS-CoV). In the present study, we used a murine monoclonal antibody specific for SARS-CoV nucleoprotein, and probes specific for a SARS-CoV RNA polymerase gene fragment, for immunohistochem. and in situ hybridization, resp., to detect SARS-CoV systematically in tissues from patients who died of SARS. SARS-CoV was found in lung, trachea/bronchus, stomach, small intestine, distal convoluted renal tubule, sweat gland, parathyroid, pituitary, pancreas,

adrenal gland, liver and cerebrum, but was not detected in esophagus, spleen, lymph node, bone marrow, heart, aorta, cerebellum, thyroid, testis, ovary, uterus or muscle. These results suggest that, in addn. to the respiratory system, the gastrointestinal tract and other organs with detectable SARS-CoV may also be targets of SARS-CoV infection. The pathol. changes in these organs may be caused directly by the cytopathic effect mediated by local replication of the SARS-CoV; or indirectly as a result of systemic responses to respiratory failure or the harmful immune response induced by viral infection. In addn. to viral spread through a respiratory route, SARS-CoV in the intestinal tract, kidney and sweat glands may be excreted via feces, urine and sweat, thereby leading to virus transmission. This study provides important information for understanding the pathogenesis of SARS-CoV infection and sheds light on possible virus transmission pathways. This data will be useful for designing new strategies for prevention and treatment of SARS.

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L21 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:539287 CAPLUS
 DOCUMENT NUMBER: 141:275951
 TITLE: Development and characterisation of neutralising monoclonal antibody to the SARS-coronavirus
 AUTHOR(S): Berry, Jody D.; Jones, Steven; Drebot, Michael A.; Andonov, Anton; Sabara, Marta; Yuan, Xin Y.; Weingartl, Hana; Fernando, Lisa; Marszal, Peter; Gren, Jason; Nicolas, Brigitte; Andonova, Maya; Ranada, Francesca; Gubbins, Michael J.; Ball, T. Blake; Kitching, Paul; Li, Yan; Kabani, Amin; Plummer, Frank
 CORPORATE SOURCE: Department of Medical Microbiology, National Centre for Foreign Animal Disease, CFIA, University of Manitoba, Winnipeg, Can.
 SOURCE: Journal of Virological Methods (2004), 120(1), 87-96
 CODEN: JMVMDH; ISSN: 0166-0934
 PUBLISHER: Elsevier Science B.V.
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB There is a global need to elucidate protective antigens expressed by the SARS-coronavirus (SARS-CoV). Monoclonal antibody reagents that recognize specific antigens on SARS-CoV are needed urgently. In this report, the development and immunochem. characterization of a panel of murine monoclonal antibodies (mAbs) against the SARS-CoV is presented, based upon their specificity, binding requirements, and biol. activity. Initial screening by ELISA, using highly purified virus as the coating antigen, resulted in the selection of 103 mAbs to the SARS virus. Subsequent screening steps reduced this panel to seventeen IgG mAbs. A single mAb, F26G15, is specific for the nucleoprotein as seen in Western immunoblot while five other mAbs react with the Spike protein. Two of these Spike-specific mAbs demonstrate the ability to neutralize SARS-CoV in vitro while another four Western immunoblot-neg. mAbs also neutralize the virus. The utility of these mAbs for diagnostic development is demonstrated. Antibody from convalescent SARS patients, but not normal human serum, is also shown to specifically compete off binding of mAbs to whole SARS-CoV. These studies highlight the importance of using standardized assays and reagents. These mAbs will be useful for the development of diagnostic tests, studies of SARS-CoV pathogenesis and vaccine development.

REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> 5 L20 IBIB ABS 1-16

L20 ANSWER 1 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2007:147094 CAPLUS
 DOCUMENT NUMBER: 147:46687
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 AUTHOR(S): Che, Xiaoyan; Liao, Zhiyong; Wang, Yadi; Qiu, Liwen; Wen, Kun; Pan, Yuxian; Xu, Hua; Mei, Yabo; Hao, Wei; Ding, Yanqing
 CORPORATE SOURCE: Zhujiang Hospital, Southern Medical University, Guangzhou, 510282, Peop. Rep. China
 SOURCE: Zhonghua Weishengwuxue He Mianyixue Zazhi (2005), 25(9), 711-715
 CODEN: ZWMZDP; ISSN: 0254-5101
 PUBLISHER: Beijing Shengwu Zhipin Yanjiuso
 DOCUMENT TYPE: Journal
 LANGUAGE: Chinese

AB A recombinant nucleocapsid (N) protein of SARS-CoV, HCoV-229E and HCoV-OC43, was obtained resp. to study antigenic relationships of N proteins between SARS-CoV and human coronaviruses 229E and OC43. The genes encoding the full-length of N proteins from SARS-CoV, HCoV-229E and HCoV-OC43 were amplified by RT-PCR and cloned into the prokaryotic expression vector pQE30. The His6-tagged N proteins were expressed in the M15 strain and further purified with affinity chromatog. The antigenicity of N proteins was analyzed by Western blot and immunofluorescence assay. The N genes of 1281, 1182 and 1359 bp from SARS-CoV, HCoV-229E and HCoV-OC43, resp. were amplified with their corresponding primer pairs. The recombinant plasmids were sequenced, and they were all in frame with sequences matching those for the N genes of the three coronaviruses. The expressed recombinant His6-tagged N proteins were identified by Western blot assay with anti-His tag monoclonal antibody. The immunoreactive protein bands with expected sizes were 47 kDa, 44 kDa and 50 kDa from SARS-CoV, HCoV-229E and HCoV-OC43, resp. The nucleocapsid proteins of SARS-CoV, HCoV-229E and HCoV-OC43 strongly and specifically reacted with the virus specific rabbit serum and with the nucleoprotein specific murine serum. No cross-reactivity was found among the nucleocapsid proteins of SARS-CoV, HCoV-229E and HCoV-OC43. The immunogenic nucleocapsid recombinant proteins from SARS-CoV, HCoV-229E and HCoV-OC43 were obtained. There was no antigenic relationship among the three N proteins.

L20 ANSWER 2 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



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 DOCUMENT NUMBER: 146:137159
 TITLE: Coronavirus nucleocapsid protein is an RNA chaperone
 AUTHOR(S): Zuniga, Sonia; Sola, Isabel; Moreno, Jose L.; Sabella, Patricia; Plana-Duran, Juan; Enjuanes, Luis
 CORPORATE SOURCE: Centro Nacional de Biotecnologia, CSIC, Department of Molecular and Cell Biology, Campus Universidad Autonoma, Madrid, 28049, Spain
 SOURCE: Virology (2007), 357(2), 215-227
 CODEN: VIRLAX; ISSN: 0042-6822

PUBLISHER: Elsevier
DOCUMENT TYPE: Journal
LANGUAGE: English

AB RNA chaperones are nonspecific nucleic acid binding proteins with long disordered regions that help RNA mols. to adopt its functional conformation. Coronavirus nucleoproteins (N) are nonspecific RNA-binding proteins with long disordered regions. Therefore, we investigated whether transmissible gastroenteritis coronavirus (TGEV) N protein was an RNA chaperone. Purified N protein enhanced hammerhead ribozyme self-cleavage and nucleic acids annealing, which are properties that define RNA chaperones. In contrast, another RNA-binding protein, PTB, did not show these activities. N protein chaperone activity was blocked by specific monoclonal antibodies. Therefore, it was concluded that TGEV N protein is an RNA chaperone. In addn., we have shown that purified severe acute respiratory syndrome (SARS)-CoV N protein also has RNA chaperone activity. In silico predictions of disordered domains showed a similar pattern for all coronavirus N proteins evaluated. Altogether, these data led us to suggest that all coronavirus N proteins might be RNA chaperones.

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L20 ANSWER 3 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:1139612 CAPLUS
DOCUMENT NUMBER: 146:140843
TITLE: Antigenic and cellular localisation analysis of the severe acute respiratory syndrome coronavirus nucleocapsid protein using monoclonal antibodies
AUTHOR(S): Bussmann, Bianca M.; Reiche, Sven; Jacob, Lotta H.; Braun, Jan Matthias; Jassoy, Christian
CORPORATE SOURCE: Institute of Virology, University of Leipzig, Leipzig, 04103, Germany
SOURCE: Virus Research (2006), 122(1-2), 119-126
CODEN: VIREDF; ISSN: 0168-1702
PUBLISHER: Elsevier B.V.
DOCUMENT TYPE: Journal
LANGUAGE: English

AB A member of the family of coronaviruses has previously been identified as the cause of the severe acute respiratory syndrome (SARS). In this study, several monoclonal antibodies against the nucleocapsid protein have been generated to examine distribution of the nucleocapsid in virus-infected cells and to study antigenic regions of the protein. Confocal microscopic anal. identified nucleocapsids packaged in vesicles in the perinuclear area indicating viral synthesis at the endoplasmic reticulum and Golgi app. The monoclonal antibodies bound to the central and carboxyterminal half of the nucleocapsid protein indicating prominent exposure and immunogenicity of this part of the protein. Antibodies recognized both linear and conformational epitopes. Predictions of antigenicity using amt.. modeling based on hydrophobicity anal. of SARS nucleoprotein could not be confirmed fully. Antibody binding to discontinuous peptides provides evidence that amino acids 274-283 and 373-382 assemble to a structural unit particularly rich in basic amino acids. In addn., amino acids 286-295, 316-325 and 361-367 that represent the epitope recognized by monoclonal antibody 6D11C1 converge indicating a well-structured C-terminal region of the SARS virus nucleocapsid protein and functional relation of the peptide regions involved. Alternatively, dimerization of the nucleocapsid protein may result in juxtaposition of the amino acid sequences 316-325 and 361-367 on one nucleoprotein mol.

to amino acid 286-295 on the second peptide. The monoclonal antibodies will be available to assess antigenicity and immunol. variabilities between different SARS CoV strains.

REFERENCE COUNT: 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 4 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:236188 CAPLUS
DOCUMENT NUMBER: 145:185561
TITLE: Time course and cellular localization of SARS-CoV nucleoprotein and RNA in lungs from fatal cases of SARS
AUTHOR(S): Nicholls, John M.; Butany, Jagdish; Poon, Leo L. M.; Chan, Kwok H.; Beh, Swan Lip; Poutanen, Susan; Peiris, J. S. Malik; Wong, Maria
CORPORATE SOURCE: Department of Pathology, The University of Hong Kong, Pok Fu Lam, Hong Kong SAR, Peop. Rep. China
SOURCE: PLoS Medicine (2006), 3(2), 222-229
CODEN: PMLEAC; ISSN: 1549-1277
URL: http://medicine.plosjournals.org/archive/1549-1676/3/2/pdf/10.1371_1549-1676_3_2_complete.pdf
PUBLISHER: Public Library of Science
DOCUMENT TYPE: Journal; (online computer file)
LANGUAGE: English

AB Background: Cellular localization of severe acute respiratory syndrome coronavirus (SARS-CoV) in the lungs of patients with SARS is important in confirming the etiol. assocn. of the virus with disease as well as in understanding the pathogenesis of the disease. To our knowledge, there have been no comprehensive studies investigating viral infection at the cellular level in humans. Methods and Findings: We collected the largest series of fatal cases of SARS with autopsy material to date by merging the pathol. material from two regions involved in the 2003 worldwide SARS outbreak in Hong Kong, China, and Toronto, Canada. We developed a monoclonal antibody against the SARS-CoV nucleoprotein and used it together with in situ hybridization (ISH) to analyze the autopsy lung tissues of 32 patients with SARS from Hong Kong and Toronto. We compared the results of these assays with the pulmonary pathologies and the clin. course of illness for each patient. SARS-CoV nucleoprotein and RNA were detected by immunohistochem. and ISH, resp., primarily in alveolar pneumocytes and, less frequently, in macrophages. Such localization was detected in four of the seven patients who died within two weeks of illness onset, and in none of the 25 patients who died later than two weeks after symptom onset. Conclusions: The pulmonary alveolar epithelium is the chief target of SARS-CoV, with macrophages infected subsequently. Viral replication appears to be limited to the first two weeks after symptom onset, with little evidence of continued widespread replication after this period. If antiviral therapy is considered for future treatment, it should be focused on this two-week period of acute clin. disease.

REFERENCE COUNT: 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 5 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2005:562710 CAPLUS
DOCUMENT NUMBER: 143:246447
TITLE: Use of monoclonal antibodies in blocking ELISA

detection of transmissible gastroenteritis virus in faeces of piglets

AUTHOR(S): Rodak, L.; Smid, B.; Nevorankova, Z.; Valicek, L.; Smitalova, R.

CORPORATE SOURCE: Veterinary Research Institute, Brno, Czech Rep.

SOURCE: Journal of Veterinary Medicine, Series B (2005), 52(3), 105-111
CODEN: JVMBE9; ISSN: 0931-1793

PUBLISHER: Blackwell Verlag GmbH

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Monoclonal antibodies (mAb) to the transmissible gastroenteritis virus (TGEV) nucleoprotein (N) and membrane protein (M) were prepd. and used for the comparative assessment of three blocking ELISA variants to detect TGEV. The competitive blocking ELISA format showed the highest sensitivity, allowing detection of 103 TCID50 TGEV/mL in culture medium. Ninety-nine porcine field fecal samples obtained from 37 herds affected with diarrhea were examd., and various TGEV levels were found in nine samples from six herds. However, only in three samples were significant TGEV concns. demonstrated. The relationship between incidence of TGEV gastroenteritis and the spread of porcine respiratory coronavirus infection in pig farms is discussed.

REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 6 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2005:409560 CAPLUS

DOCUMENT NUMBER: 142:462283

TITLE: Monoclonal antibodies specific to SARS virus nucleoprotein for immunodiagnosis of SARS

INVENTOR(S): Uchida, Yoshiaki; Fujii, Nobuyuki; Kurano, Yoshihiro; Okada, Masahisa; Kogaki, Hiroyuki; Kido, Yasuji; Miyake, Kazushige

PATENT ASSIGNEE(S): Fujirebio Inc., Japan

SOURCE: PCT Int. Appl., 41 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005042579	A1	20050512	WO 2004-JP16099	20041029
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CN 1902230	A	20070124	CN 2004-80039648	20041029
IN 2006KN01457	A	20070504	IN 2006--KN1457	20060530
US 20080254440	A1	20081016	US 2007-577310	20070222

PRIORITY APPLN. INFO.:

JP 2003-373779 A 20031031
 JP 2004-34268 A 20040210
 WO 2004-JP16099 W 20041029

AB Provided are monoclonal antibodies specific to SARS virus nucleoprotein and hybridomas producing them. These monoclonal antibodies are labeled with enzyme and used for immunodiagnosis of SARS.

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 7 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:872877 CAPLUS
 DOCUMENT NUMBER: 141:378847
 TITLE: Endogenous host elements or viral-based sequence elements for diagnosis, prognosis and therapy of viral infection, autoimmune disease and lymphoproliferative disease
 INVENTOR(S): Hu, Yu-wen; Brown, Earl
 PATENT ASSIGNEE(S): Canadian Blood Services, Can.
 SOURCE: PCT Int. Appl., 174 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004090544	A2	20041021	WO 2004-CA544	20040413
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GM, GH, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2522067	A1	20041021	CA 2004-2522067	20040413
EP 1625402	A2	20060215	EP 2004-726942	20040413
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR				
US 20060115875	A1	20060601	US 2005-248008	20051011
<u>PRIORITY APPLN. INFO.:</u>			US 2003-461137P	P 20030409
			US 2003-506779P	P 20030930
			WO 2004-CA544	W 20040413

AB A method of detecting, characterizing and treating viral infection, autoimmune disease and lymphoproliferative disease is provided. In particular, a strategy of mol. mimicry is provided for characterizing viral behavior and/or a predisposition for a given viral outcome in vivo. Novel compns. are also provided for detecting, characterizing and treating viral infections. The viral infection is caused by HCV, HIV, HTLV-1, HTLV-2, SARS-CoV, or a member of Retroviridae, Flaviviridae, Herpesviridae, Papillomaviridae, Poxviridae or Coronaviridae. The viral-based sequence element is e.g. an element of S protein sequence of an ORF1a protein sequence of SARS-CoV; a Gag, Pol or Env polyprotein of HTLV-1; a NS5A and E2 protein of HCV; bacterial virulence factor; human

endogenous retrovirus element; Peyer's patches virulence factor gipA; or an Ig selected from IgG, IgA, IgM, IgD or IgE. The treatment regime includes an anti-viral monoclonal or polyclonal antibody, or a compd. capable of binding epitope of the endogenous host element.

L20 ANSWER 8 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:541777 CAPLUS
 DOCUMENT NUMBER: 141:222968
 TITLE: Organ distribution of severe acute respiratory syndrome (SARS) associated coronavirus (SARS-CoV) in SARS patients: implications for pathogenesis and virus transmission pathways
 AUTHOR(S): Ding, Yanqing; He, Li; Zhang, Qingling; Huang, Zhongxi; Che, Xiaoyan; Hou, Jinlin; Wang, Huijun; Shen, Hong; Qiu, Liwen; Li, Zhuguo; Geng, Jian; Cai, Junjie; Han, Huixia; Li, Xin; Kang, Wei; Weng, Desheng; Liang, Ping; Jiang, Shibo
 CORPORATE SOURCE: Department of Pathology, Nan Fang Hospital, First Military Medical University, Guangzhou, Peop. Rep. China
 SOURCE: Journal of Pathology (2004), 203(2), 622-630
 CODEN: JPTLAS; ISSN: 0022-3417
 PUBLISHER: John Wiley & Sons Ltd.
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB We previously identified the major pathol. changes in the respiratory and immune systems of patients who died of severe acute respiratory syndrome (SARS) but gained little information on the organ distribution of SARS-assocd. coronavirus (SARS-CoV). In the present study, we used a murine monoclonal antibody specific for SARS-CoV nucleoprotein, and probes specific for a SARS-CoV RNA polymerase gene fragment, for immunohistochem. and in situ hybridization, resp., to detect SARS-CoV systematically in tissues from patients who died of SARS. SARS-CoV was found in lung, trachea/bronchus, stomach, small intestine, distal convoluted renal tubule, sweat gland, parathyroid, pituitary, pancreas, adrenal gland, liver and cerebrum, but was not detected in esophagus, spleen, lymph node, bone marrow, heart, aorta, cerebellum, thyroid, testis, ovary, uterus or muscle. These results suggest that, in addn. to the respiratory system, the gastrointestinal tract and other organs with detectable SARS-CoV may also be targets of SARS-CoV infection. The pathol. changes in these organs may be caused directly by the cytopathic effect mediated by local replication of the SARS-CoV; or indirectly as a result of systemic responses to respiratory failure or the harmful immune response induced by viral infection. In addn. to viral spread through a respiratory route, SARS-CoV in the intestinal tract, kidney and sweat glands may be excreted via feces, urine and sweat, thereby leading to virus transmission. This study provides important information for understanding the pathogenesis of SARS-CoV infection and sheds light on possible virus transmission pathways. This data will be useful for designing new strategies for prevention and treatment of SARS.

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 9 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:539287 CAPLUS
 DOCUMENT NUMBER: 141:275951

TITLE: Development and characterisation of neutralising monoclonal antibody to the SARS-coronavirus

AUTHOR(S): Berry, Jody D.; Jones, Steven; Drebot, Michael A.; Andonov, Anton; Sabara, Marta; Yuan, Xin Y.; Weingartl, Hana; Fernando, Lisa; Marszal, Peter; Gren, Jason; Nicolas, Brigitte; Andonova, Maya; Ranada, Francesca; Gubbins, Michael J.; Ball, T. Blake; Kitching, Paul; Li, Yan; Kabani, Amin; Plummer, Frank

CORPORATE SOURCE: Department of Medical Microbiology, National Centre for Foreign Animal Disease, CFIA, University of Manitoba, Winnipeg, Can.

SOURCE: Journal of Virological Methods (2004), 120(1), 87-96
CODEN: JVMEHD; ISSN: 0166-0934

PUBLISHER: Elsevier Science B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB There is a global need to elucidate protective antigens expressed by the SARS-coronavirus (SARS-CoV). Monoclonal antibody reagents that recognize specific antigens on SARS-CoV are needed urgently. In this report, the development and immunochem. characterization of a panel of murine monoclonal antibodies (mAbs) against the SARS-CoV is presented, based upon their specificity, binding requirements, and biol. activity. Initial screening by ELISA, using highly purified virus as the coating antigen, resulted in the selection of 103 mAbs to the SARS virus. Subsequent screening steps reduced this panel to seventeen IgG mAbs. A single mAb, F26G15, is specific for the nucleoprotein as seen in Western immunoblot while five other mAbs react with the Spike protein. Two of these Spike-specific mAbs demonstrate the ability to neutralize SARS-CoV in vitro while another four Western immunoblot-neg. mAbs also neutralize the virus. The utility of these mAbs for diagnostic development is demonstrated. Antibody from convalescent SARS patients, but not normal human serum, is also shown to specifically compete off binding of mAbs to whole SARS-CoV. These studies highlight the importance of using standardized assays and reagents. These mAbs will be useful for the development of diagnostic tests, studies of SARS-CoV pathogenesis and vaccine development.

REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 10 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2001:48484 CAPLUS

DOCUMENT NUMBER: 134:219537

TITLE: The membrane M protein carboxy terminus binds to transmissible gastroenteritis coronavirus core and contributes to core stability

AUTHOR(S): Escors, David; Ortego, Javier; Laude, Hubert; Enjuanes, Luis

CORPORATE SOURCE: Department of Molecular and Cell Biology, Centro Nacional de Biotecnologia, CSIC, Madrid, 28049, Spain

SOURCE: Journal of Virology (2001), 75(3), 1312-1324
CODEN: JOVIAM; ISSN: 0022-538X

PUBLISHER: American Society for Microbiology

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The architecture of transmissible gastroenteritis coronavirus includes three different structural levels, the envelope, an internal core, and the nucleocapsid that is released when the core is disrupted. Starting from purified virions, core structures have been reproducibly isolated as

independent entities. The cores were stabilized at basic pH and by the presence of divalent cations, with Mg²⁺ ions more effectively contributing to core stability. Core structures showed high resistance to different concns. of detergents, reducing agents, and urea and low concns. of monovalent ions (<200 mM). Cores were composed of the nucleoprotein, RNA, and the C domain of the membrane (M) protein. At high salt concns. (200 to 300 mM), the M protein was no longer assocd. with the nucleocapsid, which resulted in destruction of the core structure. A specific ionic interaction between the M protein carboxy terminus and the nucleocapsid was demonstrated using three complementary approaches: (i) a binding assay performed between a collection of M protein amino acid substitution or deletion mutants and purified nucleocapsids that led to the identification of a 16-amino-acid (aa) domain (aa 237 to 252) as being responsible for binding the M protein to the nucleocapsid; (ii) the specific inhibition of this binding by monoclonal antibodies (MAbs) binding to a carboxy-terminal M protein domain close to the indicated peptide but not by MAbs specific for the M protein amino terminus; and (iii) a 26-residue peptide, including the predicted sequence (aa 237 to 252), which specifically inhibited the binding. Direct binding of the M protein to the nucleoprotein was predicted, since degrdn. of the exposed RNA by RNase treatment did not affect the binding. It is proposed that the M protein is embedded within the virus membrane and that the C region, exposed to the interior face of the virion in a population of these mols., interacts with the nucleocapsid to which it is anchored, forming the core. Only the C region of the M protein is part of the core.

REFERENCE COUNT: 50 THERE ARE 50 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 11 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1999:336041 CAPLUS
DOCUMENT NUMBER: 131:156705
TITLE: Production, characterization, and uses of monoclonal antibodies against recombinant nucleoprotein of elk coronavirus
AUTHOR(S): Daginakatte, Girish C.; Chard-Bergstrom, Cindy; Andrews, Gordon A.; Kapil, Sanjay
CORPORATE SOURCE: Department of Diagnostic Medicine-Pathobiology, College of Veterinary Medicine, Manhattan, KS, 66506, USA
SOURCE: Clinical and Diagnostic Laboratory Immunology (1999), 6(3), 341-344
CODEN: CDIMEN; ISSN: 1071-412X
PUBLISHER: American Society for Microbiology
DOCUMENT TYPE: Journal
LANGUAGE: English
AB This is the first report of the prodn. of monoclonal antibodies against elk coronavirus. The nucleoprotein gene of elk coronavirus was amplified by PCR and was cloned and expressed in a prokaryotic expression vector. Recombinant nucleocapsid protein was used to immunize mice for the prodn. of hybridomas. Twelve hybridomas that produced monoclonal antibodies against the nucleocapsid protein of elk coronavirus were selected by an indirect fluorescent-antibody test, an ELISA, and a Western blot assay. Ten of the monoclonal antibodies were of the IgG1 isotype, one was IgG2a, and one was IgM. All had kappa light chains. By immunohistochem. four monoclonal antibodies detected bovine coronavirus and elk coronavirus in formalin-fixed intestinal tissues. Anti-nucleoprotein monoclonal antibodies were better at ruminant coronavirus detection than the anti-spike protein monoclonal

antibodies. Because nucleoprotein is a more abundant antigen than spike protein in infected cells, this was not an unexpected finding.

REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 12 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1997:723088 CAPLUS
 DOCUMENT NUMBER: 128:58017
 ORIGINAL REFERENCE NO.: 128:11239a,11242a
 TITLE: Isolation and characterization of a coronavirus from elk calves with diarrhea
 AUTHOR(S): Majhdi, F.; Minocha, H. C.; Kapil, S.
 CORPORATE SOURCE: Department of Diagnostic Medicine/Pathobiology, College of Veterinary Medicine, Kansas State University, Manhattan, KS, 66506, USA
 SOURCE: Journal of Clinical Microbiology (1997), 35(11), 2937-2942
 CODEN: JCMIDW; ISSN: 0095-1137
 PUBLISHER: American Society for Microbiology
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB This is the first report of the isolation of a coronavirus from elk calves. Two fecal samples from elk calves with diarrhea were shown to be pos. for coronavirus-like particles by electron microscopy, and the particles were propagated in the human rectal tumor-18 cell line. After 24 h, syncytia were obsd., and cell culture supernatants from both samples showed hemagglutinating activity with mouse erythrocytes. Cells infected with both elk coronavirus (ECV) isolates reacted with Z3A5, a monoclonal antibody against the spike protein of bovine coronavirus (BCV), on an indirect fluorescent antibody test. The protein profiles of both ECV isolates were similar to that of BCV as detd. by sodium dodecyl sulfate-polyacrylamide gel electrophoresis anal. On Northern blot anal., the transcriptional pattern of ECV was typical of coronaviruses, with a nested set of transcripts with common 3' end sequences. Based on a published nucleoprotein gene sequence for BCV (Mebus isolate), we arbitrarily designed two primers for amplification by PCR. After cloning, the nucleoprotein was sequenced and a high degree of homol. (99%) between the nucleoprotein gene sequences of ECV and BCV was obsd. Thus, ECV is closely related genetically and antigenically to BCV and will be a new member of antigenic group 2 of the mammalian coronaviruses, which possess hemagglutinin-esterase protein.

L20 ANSWER 13 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1995:945210 CAPLUS
 DOCUMENT NUMBER: 124:47424
 ORIGINAL REFERENCE NO.: 124:8827a,8830a
 TITLE: Experimental evidence of recombination in coronavirus infectious bronchitis virus
 AUTHOR(S): Kottier, Sanneke A.; Cavanagh, David; Britton, Paul
 CORPORATE SOURCE: Division Molecular Biology, Institute Animal Health, Compton, Newbury, Berkshire, RG20 7NN, UK
 SOURCE: Virology (1995), 213(2), 569-80
 CODEN: VIRLAX; ISSN: 0042-6822
 PUBLISHER: Academic
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB Embryonated eggs were coinfectd with two strains of the coronavirus avian infectious bronchitis virus (IBV), IBV-Beaudette and IBV-M41, to investigate whether recombination between the two strains would occur. Virions were isolated from the allantoic fluid of the coinfectd eggs and putative hybrid RNAs were detected by polymerase chain reaction (PCR), using strain-specific oligonucleotides. PCR products, of the expected sizes, were obtained as predicted from potential recombination events between the nucleoprotein (N) gene and the 3'-untranslated region of the two IBV genomes. Sequencing confirmed that they corresponded to hybrid RNAs. Virus produced as a result of the mixed infection was treated with an M41-specific neutralizing monoclonal and passaged in Vero cells, in which IBV-beaudette, but not IBV-M41, replicated. Hybrid RNA was still detectable after three serial passages. Since no IBV-M41 was detectable this confirmed that infectious recombinant genomes had been produced in the embryonated eggs. These findings not only support the circumstantial evidence, from sequencing studies of IBV field strains, that recombination occurs during replication of IBV and contributes to the diversity of IBV, but also show that coronavirus RNA recombination is not limited to mouse hepatitis virus.

L20 ANSWER 14 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1991:533524 CAPLUS
 DOCUMENT NUMBER: 115:133524
 ORIGINAL REFERENCE NO.: 115:22845a,22848a
 TITLE: Comparison of bovine coronavirus (BCV) antigens: monoclonal antibodies to the spike glycoprotein distinguish between vaccine and wild-type strains
 AUTHOR(S): Hussain, Khalid A.; Storz, Johannes; Kousoulas, Konstantin G.
 CORPORATE SOURCE: Sch. Vet. Med., Louisiana State Univ., Baton Rouge, LA, 70803, USA
 SOURCE: Virology (1991), 183(1), 442-5
 CODEN: VIRLAX; ISSN: 0042-6822
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB Monoclonal antibodies (MAbs) against two major structural proteins of the cell-adapted Mebus strain of bovine coronavirus (BCV-L9) were produced and characterized. Seven MAbs reacted with the peplomeric glycoprotein, gp100/S, while three MAbs reacted with the nucleoprotein p53/N in Western blot anal. of BCV polypeptides. MAbs to gp100/S reacted with discontinuous epitopes of gp100/S in Westerns under mild but not under std. denaturing conditions. In contrast, MAbs to p53/N reacted in both types of Westerns, and those epitopes were thus continuous. MAbs to p53/N failed to neutralize BCV infectivity, while 4 MAbs to gp100/S neutralized BCV effectively. Cross reactivity of MAbs to gp100/S specified by five virulent wild-type strains and two high passage, cell-culture-adapted strains in mildly denaturing Westerns and neutralization assays indicated that two epitopes were conserved in all seven strains, while two epitopes of the avirulent strains were not detected in the wild-type strains. Non-neutralizing MAbs of gp100/S reacted with all seven strains in Westerns with the exception of one MAb that was specific for the highly cell-adapted strain BCV-L9.

L20 ANSWER 15 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1988:470029 CAPLUS
 DOCUMENT NUMBER: 109:70029

ORIGINAL REFERENCE NO.: 109:11669a,11672a
 TITLE: Antigenic differentiation between transmissible gastroenteritis virus of swine and a related porcine respiratory coronavirus
 AUTHOR(S): Callebaut, P.; Correa, I.; Pensaert, M.; Jimenez, G.; Enjuanes, L.
 CORPORATE SOURCE: Fac. Vet. Med., State Univ. Gent, Ghent, B-9000, Belg.
 SOURCE: Journal of General Virology (1988), 69(7), 1725-30
 CODEN: JGVIAI; ISSN: 0022-1317
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB The antigenic relationship between isolated porcine respiratory coronavirus (TLM 83) and transmissible gastroenteritis (TGE) virus of swine was studied by neutralization, immunoblotting, and RIA, using TGE virus-specific monoclonal antibodies (MAbs) and polyclonal antibodies specific for both viruses. A complete two-way neutralization activity between the two viruses was found. Immunoblotting revealed cross-reactions between TLM 83 and TGE virus antigens at the level of the envelope protein (E1), the nucleoprotein (N), and the peplomer protein (E2). By virus neutralization assays and RIA with TGE virus-specific MAbs, the presence of similar epitopes in the E1 and N proteins and in the neutralization-mediating antigenic site of the E2 protein were demonstrated. E2 protein-specific MAbs, without neutralizing activity and reacting with antigenic sites B, C, and D (previously defined), failed to recognize TLM 83. These results indicated a close antigenic relationship and structural similarity between TLM 83 and TGE viruses and also suggested potential ways of differentiating between the two viruses.

L20 ANSWER 16 OF 16 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1984:3195 CAPLUS
 DOCUMENT NUMBER: 100:3195
 ORIGINAL REFERENCE NO.: 100:551a,554a
 TITLE: Synthesis and subcellular localization of the murine coronavirus nucleocapsid protein
 AUTHOR(S): Stohlgman, Stephen A.; Fleming, John O.; Patton, Chris D.; Lai, Michael M. C.
 CORPORATE SOURCE: Sch. Med., Univ. South. California, Los Angeles, CA, 90033, USA
 SOURCE: Virology (1983), 130(2), 527-32
 CODEN: VIRLAX; ISSN: 0042-6822
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB The synthesis and processing of the nucleocapsid protein (pp60) of the JHM strain of murine coronaviruses were examd. Pulse-chase expts. showed that pp60 was synthesized initially as a protein of mol. wt. ~57,000 (p57). Immunopptn. using mouse anti-JHNV antiserum indicated that p57 was virus specific. Immunopptn. with monoclonal antibodies specific for pp60 showed that p57 was antigenically related to pp60 and was not phosphorylated, whereas the intracellular protein that comigrated with the virion nucleocapsid protein, pp60, was phosphorylated. The p57 was found exclusively in the cytosol whereas the majority of pp60 was assocd. with the membrane fraction but pp60 was not an integral membrane protein.

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L16 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2007:147094 CAPLUS
 DOCUMENT NUMBER: 147:46687
 TITLE: Cloning, expressing and antigenicity analysis of nucleocapsid proteins of SARS-CoV, HCoV-229E and OC43
 AUTHOR(S): Che, Xiaoyan; Liao, Zhiyong; Wang, Yadi; Qiu, Liwen; Wen, Kun; Pan, Yuxian; Xu, Hua; Mei, Yabo; Hao, Wei; Ding, Yanqing
 CORPORATE SOURCE: Zhujiang Hospital, Southern Medical University, Guangzhou, 510282, Peop. Rep. China
 SOURCE: Zhonghua Weishengwuxue He Mianyixue Zazhi (2005), 25(9), 711-715
 CODEN: ZWMZDP; ISSN: 0254-5101
 PUBLISHER: Beijing Shengwu Zhipin Yanjiuso
 DOCUMENT TYPE: Journal
 LANGUAGE: Chinese

AB A recombinant nucleocapsid (N) protein of SARS-CoV, HCoV-229E and HCoV-OC43, was obtained resp. to study antigenic relationships of N proteins between SARS-CoV and human coronaviruses 229E and OC43. The genes encoding the full-length of N proteins from SARS-CoV, HCoV-229E and HCoV-OC43 were amplified by RT-PCR and cloned into the prokaryotic expression vector pQE30. The His6-tagged N proteins were expressed in the M15 strain and further purified with affinity chromatog. The antigenicity of N proteins was analyzed by Western blot and immunofluorescence assay. The N genes of 1281, 1182 and 1359 bp from SARS-CoV, HCoV-229E and HCoV-OC43, resp. were amplified with their corresponding primer pairs. The recombinant plasmids were sequenced, and they were all in frame with sequences matching those for the N genes of the three coronaviruses. The expressed recombinant His6-tagged N proteins were identified by Western blot assay with anti-His tag monoclonal antibody. The immunoreactive protein bands with expected sizes were 47 kDa, 44 kDa and 50 kDa from SARS-CoV, HCoV-229E and HCoV-OC43, resp. The nucleocapsid proteins of SARS-CoV, HCoV-229E and HCoV-OC43 strongly and specifically reacted with the virus specific rabbit serum and with the nucleoprotein specific murine serum. No cross-reactivity was found among the nucleocapsid proteins of SARS-CoV, HCoV-229E and HCoV-OC43. The immunogenic nucleocapsid recombinant proteins from SARS-CoV, HCoV-229E and HCoV-OC43 were obtained. There was no antigenic relationship among the three N proteins.

L16 ANSWER 2 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:1263772 CAPLUS
 DOCUMENT NUMBER: 146:137159
 TITLE: Coronavirus nucleocapsid protein is an RNA chaperone
 AUTHOR(S): Zuniga, Sonia; Sola, Isabel; Moreno, Jose L.; Sabella, Patricia; Plana-Duran, Juan; Enjuanes, Luis
 CORPORATE SOURCE: Centro Nacional de Biotecnologia, CSIC, Department of Molecular and Cell Biology, Campus Universidad Autonoma, Madrid, 28049, Spain
 SOURCE: Virology (2007), 357(2), 215-227
 CODEN: VIRLAX; ISSN: 0042-6822
 PUBLISHER: Elsevier
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB RNA chaperones are nonspecific nucleic acid binding proteins with long disordered regions that help RNA mols. to adopt its functional conformation. Coronavirus nucleoproteins (N) are nonspecific

RNA-binding proteins with long disordered regions. Therefore, we investigated whether transmissible gastroenteritis coronavirus (TGEV) N protein was an RNA chaperone. Purified N protein enhanced hammerhead ribozyme self-cleavage and nucleic acids annealing, which are properties that define RNA chaperones. In contrast, another RNA-binding protein, PTB, did not show these activities. N protein chaperone activity was blocked by specific monoclonal antibodies. Therefore, it was concluded that TGEV N protein is an RNA chaperone. In addn., we have shown that purified severe acute respiratory syndrome (SARS)-CoV N protein also has RNA chaperone activity. In silico predictions of disordered domains showed a similar pattern for all coronavirus N proteins evaluated. Altogether, these data led us to suggest that all coronavirus N proteins might be RNA chaperones.

REFERENCE COUNT: 83 THERE ARE 83 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L16 ANSWER 3 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:1139612 CAPLUS
 DOCUMENT NUMBER: 146:140843
 TITLE: Antigenic and cellular localisation analysis of the severe acute respiratory syndrome coronavirus nucleocapsid protein using monoclonal antibodies
 AUTHOR(S): Bussmann, Bianca M.; Reiche, Sven; Jacob, Lotta H.; Braun, Jan Matthias; Jassoy, Christian
 CORPORATE SOURCE: Institute of Virology, University of Leipzig, Leipzig, 04103, Germany
 SOURCE: Virus Research (2006), 122(1-2), 119-126
 CODEN: VIREDF; ISSN: 0168-1702
 PUBLISHER: Elsevier B.V.
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB A member of the family of coronaviruses has previously been identified as the cause of the severe acute respiratory syndrome (SARS). In this study, several monoclonal antibodies against the nucleocapsid protein have been generated to examine distribution of the nucleocapsid in virus-infected cells and to study antigenic regions of the protein. Confocal microscopic anal. identified nucleocapsids packaged in vesicles in the perinuclear area indicating viral synthesis at the endoplasmic reticulum and Golgi app. The monoclonal antibodies bound to the central and carboxyterminal half of the nucleocapsid protein indicating prominent exposure and immunogenicity of this part of the protein. Antibodies recognized both linear and conformational epitopes. Predictions of antigenicity using amt.. modeling based on hydrophobicity anal. of SARS nucleoprotein could not be confirmed fully. Antibody binding to discontinuous peptides provides evidence that amino acids 274-283 and 373-382 assemble to a structural unit particularly rich in basic amino acids. In addn., amino acids 286-295, 316-325 and 361-367 that represent the epitope recognized by monoclonal antibody 6D11C1 converge indicating a well-structured C-terminal region of the SARS virus nucleocapsid protein and functional relation of the peptide regions involved. Alternatively, dimerization of the nucleocapsid protein may result in juxtaposition of the amino acid sequences 316-325 and 361-367 on one nucleoprotein mol. to amino acid 286-295 on the second peptide. The monoclonal antibodies will be available to assess antigenicity and immunol. variabilities between different SARS CoV strains.

REFERENCE COUNT: 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L16 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:236188 CAPLUS
 DOCUMENT NUMBER: 145:185561
 TITLE: Time course and cellular localization of SARS-CoV nucleoprotein and RNA in lungs from fatal cases of SARS
 AUTHOR(S): Nicholls, John M.; Butany, Jagdish; Poon, Leo L. M.; Chan, Kwok H.; Beh, Swan Lip; Poutanen, Susan; Peiris, J. S. Malik; Wong, Maria
 CORPORATE SOURCE: Department of Pathology, The University of Hong Kong, Pok Fu Lam, Hong Kong SAR, Peop. Rep. China
 SOURCE: PLoS Medicine (2006), 3(2), 222-229
 CODEN: PMLEAC; ISSN: 1549-1277
 URL: <http://medicine.plosjournals.org/archive/1549-1676/3/2/pdf/10.1371.1549-1676.3.2.complete.pdf>
 PUBLISHER: Public Library of Science
 DOCUMENT TYPE: Journal; (online computer file)
 LANGUAGE: English

AB Background: Cellular localization of severe acute respiratory syndrome coronavirus (SARS-CoV) in the lungs of patients with SARS is important in confirming the etiol. assocn. of the virus with disease as well as in understanding the pathogenesis of the disease. To our knowledge, there have been no comprehensive studies investigating viral infection at the cellular level in humans. Methods and Findings: We collected the largest series of fatal cases of SARS with autopsy material to date by merging the pathol. material from two regions involved in the 2003 worldwide SARS outbreak in Hong Kong, China, and Toronto, Canada. We developed a monoclonal antibody against the SARS-CoV nucleoprotein and used it together with in situ hybridization (ISH) to analyze the autopsy lung tissues of 32 patients with SARS from Hong Kong and Toronto. We compared the results of these assays with the pulmonary pathologies and the clin. course of illness for each patient. SARS-CoV nucleoprotein and RNA were detected by immunohistochem. and ISH, resp., primarily in alveolar pneumocytes and, less frequently, in macrophages. Such localization was detected in four of the seven patients who died within two weeks of illness onset, and in none of the 25 patients who died later than two weeks after symptom onset. Conclusions: The pulmonary alveolar epithelium is the chief target of SARS-CoV, with macrophages infected subsequently. Viral replication appears to be limited to the first two weeks after symptom onset, with little evidence of continued widespread replication after this period. If antiviral therapy is considered for future treatment, it should be focused on this two-week period of acute clin. disease.

REFERENCE COUNT: 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L16 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2005:409560 CAPLUS
 DOCUMENT NUMBER: 142:462283
 TITLE: Monoclonal antibodies specific to SARS virus nucleoprotein for immunodiagnosis of SARS
 INVENTOR(S): Uchida, Yoshiaki; Fujii, Nobuyuki; Kurano, Yoshihiro; Okada, Masahisa; Kogaki, Hiroyuki; Kido, Yasuji; Miyake, Kazushige
 PATENT ASSIGNEE(S): Fujirebio Inc., Japan
 SOURCE: PCT Int. Appl., 41 pp.

CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005042579	A1	20050512	WO 2004-JP16099	20041029
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CN 1902230	A	20070124	CN 2004-80039648	20041029
IN 2006KN01457	A	20070504	IN 2006-KN1457	20060530
US 20080254440	A1	20081016	US 2007-577310	20070222
PRIORITY APPLN. INFO.:			JP 2003-373779	A 20031031
			JP 2004-34268	A 20040210
			WO 2004-JP16099	W 20041029

AB Provided are monoclonal antibodies specific to SARS virus nucleoprotein and hybridomas producing them. These monoclonal antibodies are labeled with enzyme and used for immunodiagnosis of SARS.

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L16 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:872877 CAPLUS
 DOCUMENT NUMBER: 141:378847
 TITLE: Endogenous host elements or viral-based sequence elements for diagnosis, prognosis and therapy of viral infection, autoimmune disease and lymphoproliferative disease
 INVENTOR(S): Hu, Yu-wen; Brown, Earl
 PATENT ASSIGNEE(S): Canadian Blood Services, Can.
 SOURCE: PCT Int. Appl., 174 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004090544	A2	20041021	WO 2004-CA544	20040413
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,				

BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
 ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
 SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
 TD, TG

CA 2522067 A1 20041021 CA 2004-2522067 20040413

EP 1625402 A2 20060215 EP 2004-726942 20040413

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR

US 20060115875 A1 20060601 US 2005-248008 20051011

PRIORITY APPLN. INFO.: US 2003-461137P P 20030409

US 2003-506779P P 20030930

WO 2004-CA544 W 20040413

AB A method of detecting, characterizing and treating viral infection, autoimmune disease and lymphoproliferative disease is provided. In particular, a strategy of mol. mimicry is provided for characterizing viral behavior and/or a predisposition for a given viral outcome in vivo. Novel compns. are also provided for detecting, characterizing and treating viral infections. The viral infection is caused by HCV, HIV, HTLV-1, HTLV-2, SARS-CoV, or a member of Retroviridae, Flaviviridae, Herpesviridae, Papillomaviridae, Poxviridae or Coronaviridae. The viral-based sequence element is e.g. an element of S protein sequence of an ORF1a protein sequence of SARS-CoV; a Gag, Pol or Env polypeptide of HTLV-1; a NS5A and E2 protein of HCV; bacterial virulence factor; human endogenous retrovirus element; Peyer's patches virulence factor gipA; or an Ig selected from IgG, IgA, IgM, IgD or IgE. The treatment regime includes an anti-viral monoclonal or polyclonal antibody, or a compd. capable of binding epitope of the endogenous host element.

L16 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:541777 CAPLUS

DOCUMENT NUMBER: 141:222968

TITLE: Organ distribution of severe acute respiratory syndrome (SARS) associated coronavirus (SARS-CoV) in SARS patients: implications for pathogenesis and virus transmission pathways

AUTHOR(S): Ding, Yanqing; He, Li; Zhang, Qingling; Huang, Zhongxi; Che, Xiaoyan; Hou, Jinlin; Wang, Huijun; Shen, Hong; Qiu, Liwen; Li, Zhuguo; Geng, Jian; Cai, Junjie; Han, Huixia; Li, Xin; Kang, Wei; Weng, Desheng; Liang, Ping; Jiang, Shibo

CORPORATE SOURCE: Department of Pathology, Nan Fang Hospital, First Military Medical University, Guangzhou, Peop. Rep. China

SOURCE: Journal of Pathology (2004), 203(2), 622-630

CODEN: JPTLAS; ISSN: 0022-3417

PUBLISHER: John Wiley & Sons Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB We previously identified the major pathol. changes in the respiratory and immune systems of patients who died of severe acute respiratory syndrome (SARS) but gained little information on the organ distribution of SARS-assocd. coronavirus (SARS-CoV). In the present study, we used a murine monoclonal antibody specific for SARS-CoV nucleoprotein, and probes specific for a SARS-CoV RNA polymerase gene fragment, for immunohistochem. and in situ hybridization, resp., to detect SARS-CoV systematically in tissues from patients who died of SARS. SARS-CoV was found in lung, trachea/bronchus, stomach, small intestine, distal convoluted renal tubule, sweat gland, parathyroid, pituitary, pancreas,

adrenal gland, liver and cerebrum, but was not detected in esophagus, spleen, lymph node, bone marrow, heart, aorta, cerebellum, thyroid, testis, ovary, uterus or muscle. These results suggest that, in addn. to the respiratory system, the gastrointestinal tract and other organs with detectable SARS-CoV may also be targets of SARS-CoV infection. The pathol. changes in these organs may be caused directly by the cytopathic effect mediated by local replication of the SARS-CoV; or indirectly as a result of systemic responses to respiratory failure or the harmful immune response induced by viral infection. In addn. to viral spread through a respiratory route, SARS-CoV in the intestinal tract, kidney and sweat glands may be excreted via feces, urine and sweat, thereby leading to virus transmission. This study provides important information for understanding the pathogenesis of SARS-CoV infection and sheds light on possible virus transmission pathways. This data will be useful for designing new strategies for prevention and treatment of SARS.

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L16 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:539287 CAPLUS
 DOCUMENT NUMBER: 141:275951
 TITLE: Development and characterisation of neutralising monoclonal antibody to the SARS-coronavirus
 AUTHOR(S): Berry, Jody D.; Jones, Steven; Drebot, Michael A.; Andonov, Anton; Sabara, Marta; Yuan, Xin Y.; Weingartl, Hana; Fernando, Lisa; Marszal, Peter; Gren, Jason; Nicolas, Brigitte; Andonova, Maya; Ranada, Francesca; Gubbins, Michael J.; Ball, T. Blake; Kitching, Paul; Li, Yan; Kabani, Amin; Plummer, Frank
 CORPORATE SOURCE: Department of Medical Microbiology, National Centre for Foreign Animal Disease, CFIA, University of Manitoba, Winnipeg, Can.
 SOURCE: Journal of Virological Methods (2004), 120(1), 87-96
 CODEN: JVMEHD; ISSN: 0166-0934
 PUBLISHER: Elsevier Science B.V.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB There is a global need to elucidate protective antigens expressed by the SARS-coronavirus (SARS-CoV). Monoclonal antibody reagents that recognize specific antigens on SARS-CoV are needed urgently. In this report, the development and immunochem. characterization of a panel of murine monoclonal antibodies (mAbs) against the SARS-CoV is presented, based upon their specificity, binding requirements, and biol. activity. Initial screening by ELISA, using highly purified virus as the coating antigen, resulted in the selection of 103 mAbs to the SARS virus. Subsequent screening steps reduced this panel to seventeen IgG mAbs. A single mAb, F26G15, is specific for the nucleoprotein as seen in Western immunoblot while five other mAbs react with the Spike protein. Two of these Spike-specific mAbs demonstrate the ability to neutralize SARS-CoV in vitro while another four Western immunoblot-neg. mAbs also neutralize the virus. The utility of these mAbs for diagnostic development is demonstrated. Antibody from convalescent SARS patients, but not normal human serum, is also shown to specifically compete off binding of mAbs to whole SARS-CoV. These studies highlight the importance of using standardized assays and reagents. These mAbs will be useful for the development of diagnostic tests, studies of SARS-CoV pathogenesis and vaccine development.
 REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> 5 L15 IBIB ABS 1-15

L15 ANSWER 1 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2007:147094 CAPLUS
 DOCUMENT NUMBER: 147:46687
 TITLE: Cloning, expressing and antigenicity analysis of nucleocapsid proteins of SARS-CoV, HCoV-229E and OC43
 AUTHOR(S): Che, Xiaoyan; Liao, Zhiyong; Wang, Yadi; Qiu, Liwen; Wen, Kun; Pan, Yuxian; Xu, Hua; Mei, Yabo; Hao, Wei; Ding, Yanqing
 CORPORATE SOURCE: Zhujiang Hospital, Southern Medical University, Guangzhou, 510282, Peop. Rep. China
 SOURCE: Zhonghua Weishengwuxue He Mianyixue Zazhi (2005), 25(9), 711-715
 CODEN: ZWMZDP; ISSN: 0254-5101
 PUBLISHER: Beijing Shengwu Zhipin Yanjiuso
 DOCUMENT TYPE: Journal
 LANGUAGE: Chinese

AB A recombinant nucleocapsid (N) protein of SARS-CoV, HCoV-229E and HCoV-OC43, was obtained resp. to study antigenic relationships of N proteins between SARS-CoV and human coronaviruses 229E and OC43. The genes encoding the full-length of N proteins from SARS-CoV, HCoV-229E and HCoV-OC43 were amplified by RT-PCR and cloned into the prokaryotic expression vector pQE30. The His6-tagged N proteins were expressed in the M15 strain and further purified with affinity chromatog. The antigenicity of N proteins was analyzed by Western blot and immunofluorescence assay. The N genes of 1281, 1182 and 1359 bp from SARS-CoV, HCoV-229E and HCoV-OC43, resp. were amplified with their corresponding primer pairs. The recombinant plasmids were sequenced, and they were all in frame with sequences matching those for the N genes of the three coronaviruses. The expressed recombinant His6-tagged N proteins were identified by Western blot assay with anti-His tag monoclonal antibody. The immunoreactive protein bands with expected sizes were 47 kDa, 44 kDa and 50 kDa from SARS-CoV, HCoV-229E and HCoV-OC43, resp. The nucleocapsid proteins of SARS-CoV, HCoV-229E and HCoV-OC43 strongly and specifically reacted with the virus specific rabbit serum and with the nucleoprotein specific murine serum. No cross-reactivity was found among the nucleocapsid proteins of SARS-CoV, HCoV-229E and HCoV-OC43. The immunogenic nucleocapsid recombinant proteins from SARS-CoV, HCoV-229E and HCoV-OC43 were obtained. There was no antigenic relationship among the three N proteins.

L15 ANSWER 2 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:1263772 CAPLUS
 DOCUMENT NUMBER: 146:137159
 TITLE: Coronavirus nucleocapsid protein is an RNA chaperone
 AUTHOR(S): Zuniga, Sonia; Sola, Isabel; Moreno, Jose L.; Sabella, Patricia; Plana-Duran, Juan; Enjuanes, Luis
 CORPORATE SOURCE: Centro Nacional de Biotecnologia, CSIC, Department of Molecular and Cell Biology, Campus Universidad Autonoma, Madrid, 28049, Spain
 SOURCE: Virology (2007), 357(2), 215-227
 CODEN: VIRLAX; ISSN: 0042-6822

PUBLISHER: Elsevier
DOCUMENT TYPE: Journal
LANGUAGE: English

AB RNA chaperones are nonspecific nucleic acid binding proteins with long disordered regions that help RNA mols. to adopt its functional conformation. Coronavirus nucleoproteins (N) are nonspecific RNA-binding proteins with long disordered regions. Therefore, we investigated whether transmissible gastroenteritis coronavirus (TGEV) N protein was an RNA chaperone. Purified N protein enhanced hammerhead ribozyme self-cleavage and nucleic acids annealing, which are properties that define RNA chaperones. In contrast, another RNA-binding protein, PTB, did not show these activities. N protein chaperone activity was blocked by specific monoclonal antibodies. Therefore, it was concluded that TGEV N protein is an RNA chaperone. In addn., we have shown that purified severe acute respiratory syndrome (SARS)-CoV N protein also has RNA chaperone activity. In silico predictions of disordered domains showed a similar pattern for all coronavirus N proteins evaluated. Altogether, these data led us to suggest that all coronavirus N proteins might be RNA chaperones.

REFERENCE COUNT: 83 THERE ARE 83 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 3 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:1139612 CAPLUS
DOCUMENT NUMBER: 146:140843
TITLE: Antigenic and cellular localisation analysis of the severe acute respiratory syndrome coronavirus nucleocapsid protein using monoclonal antibodies
AUTHOR(S): Bussmann, Bianca M.; Reiche, Sven; Jacob, Lotta H.; Braun, Jan Matthias; Jassoy, Christian
CORPORATE SOURCE: Institute of Virology, University of Leipzig, Leipzig, 04103, Germany
SOURCE: Virus Research (2006), 122(1-2), 119-126
CODEN: VIREDF; ISSN: 0168-1702
PUBLISHER: Elsevier B.V.
DOCUMENT TYPE: Journal
LANGUAGE: English

AB A member of the family of coronaviruses has previously been identified as the cause of the severe acute respiratory syndrome (SARS). In this study, several monoclonal antibodies against the nucleocapsid protein have been generated to examine distribution of the nucleocapsid in virus-infected cells and to study antigenic regions of the protein. Confocal microscopic anal. identified nucleocapsids packaged in vesicles in the perinuclear area indicating viral synthesis at the endoplasmic reticulum and Golgi app. The monoclonal antibodies bound to the central and carboxyterminal half of the nucleocapsid protein indicating prominent exposure and immunogenicity of this part of the protein. Antibodies recognized both linear and conformational epitopes. Predictions of antigenicity using amt.. modeling based on hydrophobicity anal. of SARS nucleoprotein could not be confirmed fully. Antibody binding to discontinuous peptides provides evidence that amino acids 274-283 and 373-382 assemble to a structural unit particularly rich in basic amino acids. In addn., amino acids 286-295, 316-325 and 361-367 that represent the epitope recognized by monoclonal antibody 6D11C1 converge indicating a well-structured C-terminal region of the SARS virus nucleocapsid protein and functional relation of the peptide regions involved. Alternatively, dimerization of the nucleocapsid protein may result in juxtaposition of the amino acid sequences 316-325 and 361-367 on

one nucleoprotein mol. to amino acid 286-295 on the second peptide. The monoclonal antibodies will be available to assess antigenicity and immunol. variabilities between different SARS CoV strains.

REFERENCE COUNT: 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 4 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2006:236188 CAPLUS
DOCUMENT NUMBER: 145:185561
TITLE: Time course and cellular localization of SARS-CoV nucleoprotein and RNA in lungs from fatal cases of SARS
AUTHOR(S): Nicholls, John M.; Butany, Jagdish; Poon, Leo L. M.; Chan, Kwok H.; Beh, Swan Lip; Poutanen, Susan; Peiris, J. S. Malik; Wong, Maria
CORPORATE SOURCE: Department of Pathology, The University of Hong Kong, Pok Fu Lam, Hong Kong SAR, Peop. Rep. China
SOURCE: PLoS Medicine (2006), 3(2), 222-229
CODEN: PMLEAC; ISSN: 1549-1277
URL: http://medicine.plosjournals.org/archive/1549-1676/3/2/pdf/10.1371_1549-1676_3_2_complete.pdf
PUBLISHER: Public Library of Science
DOCUMENT TYPE: Journal; (online computer file)
LANGUAGE: English

AB Background: Cellular localization of severe acute respiratory syndrome coronavirus (SARS-CoV) in the lungs of patients with SARS is important in confirming the etiol. assocn. of the virus with disease as well as in understanding the pathogenesis of the disease. To our knowledge, there have been no comprehensive studies investigating viral infection at the cellular level in humans. Methods and Findings: We collected the largest series of fatal cases of SARS with autopsy material to date by merging the pathol. material from two regions involved in the 2003 worldwide SARS outbreak in Hong Kong, China, and Toronto, Canada. We developed a monoclonal antibody against the SARS-CoV nucleoprotein and used it together with in situ hybridization (ISH) to analyze the autopsy lung tissues of 32 patients with SARS from Hong Kong and Toronto. We compared the results of these assays with the pulmonary pathologies and the clin. course of illness for each patient. SARS-CoV nucleoprotein and RNA were detected by immunohistochem. and ISH, resp., primarily in alveolar pneumocytes and, less frequently, in macrophages. Such localization was detected in four of the seven patients who died within two weeks of illness onset, and in none of the 25 patients who died later than two weeks after symptom onset. Conclusions: The pulmonary alveolar epithelium is the chief target of SARS-CoV, with macrophages infected subsequently. Viral replication appears to be limited to the first two weeks after symptom onset, with little evidence of continued widespread replication after this period. If antiviral therapy is considered for future treatment, it should be focused on this two-week period of acute clin. disease.

REFERENCE COUNT: 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 5 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2005:562710 CAPLUS
DOCUMENT NUMBER: 143:246447
TITLE: Use of monoclonal antibodies in blocking ELISA

detection of transmissible gastroenteritis virus in faeces of piglets

AUTHOR(S): Rodak, L.; Smid, B.; Nevorankova, Z.; Valicek, L.; Smitalova, R.

CORPORATE SOURCE: Veterinary Research Institute, Brno, Czech Rep.

SOURCE: Journal of Veterinary Medicine, Series B (2005), 52(3), 105-111
CODEN: JVMBE9; ISSN: 0931-1793

PUBLISHER: Blackwell Verlag GmbH

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Monoclonal antibodies (mAb) to the transmissible gastroenteritis virus (TGEV) nucleoprotein (N) and membrane protein (M) were prepd. and used for the comparative assessment of three blocking ELISA variants to detect TGEV. The competitive blocking ELISA format showed the highest sensitivity, allowing detection of 103 TCID50 TGEV/mL in culture medium. Ninety-nine porcine field fecal samples obtained from 37 herds affected with diarrhea were examd., and various TGEV levels were found in nine samples from six herds. However, only in three samples were significant TGEV concns. demonstrated. The relationship between incidence of TGEV gastroenteritis and the spread of porcine respiratory coronavirus infection in pig farms is discussed.

REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 6 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2005:409560 CAPLUS

DOCUMENT NUMBER: 142:462283

TITLE: Monoclonal antibodies specific to SARS virus nucleoprotein for immunodiagnosis of SARS

INVENTOR(S): Uchida, Yoshiaki; Fujii, Nobuyuki; Kurano, Yoshihiro; Okada, Masahisa; Kogaki, Hiroyuki; Kido, Yasuji; Miyake, Kazushige

PATENT ASSIGNEE(S): Fujirebio Inc., Japan

SOURCE: PCT Int. Appl., 41 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
<u>WO 2005042579</u>	A1	20050512	<u>WO 2004-JP16099</u>	20041029
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
<u>CN 1902230</u>	A	20070124	<u>CN 2004-80039648</u>	20041029
<u>IN 2006KN01457</u>	A	20070504	<u>IN 2006--KN1457</u>	20060530
<u>US 20080254440</u>	A1	20081016	<u>US 2007-577310</u>	20070222

PRIORITY APPLN. INFO.:

JP 2003-373779 A 20031031
 JP 2004-34268 A 20040210
 WO 2004-JP16099 W 20041029

AB Provided are monoclonal antibodies specific to SARS virus nucleoprotein and hybridomas producing them. These monoclonal antibodies are labeled with enzyme and used for immunodiagnosis of SARS.

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 7 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:872877 CAPLUS
 DOCUMENT NUMBER: 141:378847
 TITLE: Endogenous host elements or viral-based sequence elements for diagnosis, prognosis and therapy of viral infection, autoimmune disease and lymphoproliferative disease
 INVENTOR(S): Hu, Yu-wen; Brown, Earl
 PATENT ASSIGNEE(S): Canadian Blood Services, Can.
 SOURCE: PCT Int. Appl., 174 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004090544	A2	20041021	WO 2004-CA544	20040413
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2522067	A1	20041021	CA 2004-2522067	20040413
EP 1625402	A2	20060215	EP 2004-726942	20040413
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR				
US 20060115875	A1	20060601	US 2005-248008	20051011
<u>PRIORITY APPLN. INFO.:</u>			US 2003-461137P	P 20030409
			US 2003-506779P	P 20030930
			WO 2004-CA544	W 20040413

AB A method of detecting, characterizing and treating viral infection, autoimmune disease and lymphoproliferative disease is provided. In particular, a strategy of mol. mimicry is provided for characterizing viral behavior and/or a predisposition for a given viral outcome in vivo. Novel compns. are also provided for detecting, characterizing and treating viral infections. The viral infection is caused by HCV, HIV, HTLV-1, HTLV-2, SARS-CoV, or a member of Retroviridae, Flaviviridae, Herpesviridae, Papillomaviridae, Poxviridae or Coronaviridae. The viral-based sequence element is e.g. an element of S protein sequence of an ORF1a protein sequence of SARS-CoV; a Gag, Pol or Env polyprotein of HTLV-1; a NS5A and E2 protein of HCV; bacterial virulence factor; human

endogenous retrovirus element; Peyer's patches virulence factor gipA; or an Ig selected from IgG, IgA, IgM, IgD or IgE. The treatment regime includes an anti-viral monoclonal or polyclonal antibody, or a compd. capable of binding epitope of the endogenous host element.

L15 ANSWER 8 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:541777 CAPLUS
 DOCUMENT NUMBER: 141:222968
 TITLE: Organ distribution of severe acute respiratory syndrome (SARS) associated coronavirus (SARS-CoV) in SARS patients: implications for pathogenesis and virus transmission pathways
 AUTHOR(S): Ding, Yanqing; He, Li; Zhang, Qingling; Huang, Zhongxi; Che, Xiaoyan; Hou, Jinlin; Wang, Huijun; Shen, Hong; Qiu, Liwen; Li, Zhuguo; Geng, Jian; Cai, Junjie; Han, Huixia; Li, Xin; Kang, Wei; Weng, Desheng; Liang, Ping; Jiang, Shibo
 CORPORATE SOURCE: Department of Pathology, Nan Fang Hospital, First Military Medical University, Guangzhou, Peop. Rep. China
 SOURCE: Journal of Pathology (2004), 203(2), 622-630
 CODEN: JPTLAS; ISSN: 0022-3417
 PUBLISHER: John Wiley & Sons Ltd.
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB We previously identified the major pathol. changes in the respiratory and immune systems of patients who died of severe acute respiratory syndrome (SARS) but gained little information on the organ distribution of SARS-assocd. coronavirus (SARS-CoV). In the present study, we used a murine monoclonal antibody specific for SARS-CoV nucleoprotein, and probes specific for a SARS-CoV RNA polymerase gene fragment, for immunohistochem. and in situ hybridization, resp., to detect SARS-CoV systematically in tissues from patients who died of SARS. SARS-CoV was found in lung, trachea/bronchus, stomach, small intestine, distal convoluted renal tubule, sweat gland, parathyroid, pituitary, pancreas, adrenal gland, liver and cerebrum, but was not detected in esophagus, spleen, lymph node, bone marrow, heart, aorta, cerebellum, thyroid, testis, ovary, uterus or muscle. These results suggest that, in addn. to the respiratory system, the gastrointestinal tract and other organs with detectable SARS-CoV may also be targets of SARS-CoV infection. The pathol. changes in these organs may be caused directly by the cytopathic effect mediated by local replication of the SARS-CoV; or indirectly as a result of systemic responses to respiratory failure or the harmful immune response induced by viral infection. In addn. to viral spread through a respiratory route, SARS-CoV in the intestinal tract, kidney and sweat glands may be excreted via feces, urine and sweat, thereby leading to virus transmission. This study provides important information for understanding the pathogenesis of SARS-CoV infection and sheds light on possible virus transmission pathways. This data will be useful for designing new strategies for prevention and treatment of SARS.

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 9 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2004:539287 CAPLUS
 DOCUMENT NUMBER: 141:275951

TITLE: Development and characterisation of neutralising monoclonal antibody to the SARS-coronavirus

AUTHOR(S): Berry, Jody D.; Jones, Steven; Drebot, Michael A.; Andonov, Anton; Sabara, Marta; Yuan, Xin Y.; Weingartl, Hana; Fernando, Lisa; Marszal, Peter; Gren, Jason; Nicolas, Brigitte; Andonova, Maya; Ranada, Francesca; Gubbins, Michael J.; Ball, T. Blake; Kitching, Paul; Li, Yan; Kabani, Amin; Plummer, Frank

CORPORATE SOURCE: Department of Medical Microbiology, National Centre for Foreign Animal Disease, CFIA, University of Manitoba, Winnipeg, Can.

SOURCE: Journal of Virological Methods (2004), 120(1), 87-96
CODEN: JVMEHD; ISSN: 0166-0934

PUBLISHER: Elsevier Science B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB There is a global need to elucidate protective antigens expressed by the SARS-coronavirus (SARS-CoV). Monoclonal antibody reagents that recognize specific antigens on SARS-CoV are needed urgently. In this report, the development and immunochem. characterization of a panel of murine monoclonal antibodies (mAbs) against the SARS-CoV is presented, based upon their specificity, binding requirements, and biol. activity. Initial screening by ELISA, using highly purified virus as the coating antigen, resulted in the selection of 103 mAbs to the SARS virus. Subsequent screening steps reduced this panel to seventeen IgG mAbs. A single mAb, F26G15, is specific for the nucleoprotein as seen in Western immunoblot while five other mAbs react with the Spike protein. Two of these Spike-specific mAbs demonstrate the ability to neutralize SARS-CoV in vitro while another four Western immunoblot-neg. mAbs also neutralize the virus. The utility of these mAbs for diagnostic development is demonstrated. Antibody from convalescent SARS patients, but not normal human serum, is also shown to specifically compete off binding of mAbs to whole SARS-CoV. These studies highlight the importance of using standardized assays and reagents. These mAbs will be useful for the development of diagnostic tests, studies of SARS-CoV pathogenesis and vaccine development.

REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 10 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 2001:48484 CAPLUS

DOCUMENT NUMBER: 134:219537

TITLE: The membrane M protein carboxy terminus binds to transmissible gastroenteritis coronavirus core and contributes to core stability

AUTHOR(S): Escors, David; Ortego, Javier; Laude, Hubert; Enjuanes, Luis

CORPORATE SOURCE: Department of Molecular and Cell Biology, Centro Nacional de Biotecnologia, CSIC, Madrid, 28049, Spain

SOURCE: Journal of Virology (2001), 75(3), 1312-1324
CODEN: JOVIAM; ISSN: 0022-538X

PUBLISHER: American Society for Microbiology

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The architecture of transmissible gastroenteritis coronavirus includes three different structural levels, the envelope, an internal core, and the nucleocapsid that is released when the core is disrupted. Starting from purified virions, core structures have been reproducibly isolated as

independent entities. The cores were stabilized at basic pH and by the presence of divalent cations, with Mg²⁺ ions more effectively contributing to core stability. Core structures showed high resistance to different concns. of detergents, reducing agents, and urea and low concns. of monovalent ions (<200 mM). Cores were composed of the nucleoprotein, RNA, and the C domain of the membrane (M) protein. At high salt concns. (200 to 300 mM), the M protein was no longer assocd. with the nucleocapsid, which resulted in destruction of the core structure. A specific ionic interaction between the M protein carboxy terminus and the nucleocapsid was demonstrated using three complementary approaches: (i) a binding assay performed between a collection of M protein amino acid substitution or deletion mutants and purified nucleocapsids that led to the identification of a 16-amino-acid (aa) domain (aa 237 to 252) as being responsible for binding the M protein to the nucleocapsid; (ii) the specific inhibition of this binding by monoclonal antibodies (MAbs) binding to a carboxy-terminal M protein domain close to the indicated peptide but not by MAbs specific for the M protein amino terminus; and (iii) a 26-residue peptide, including the predicted sequence (aa 237 to 252), which specifically inhibited the binding. Direct binding of the M protein to the nucleoprotein was predicted, since degrdn. of the exposed RNA by RNase treatment did not affect the binding. It is proposed that the M protein is embedded within the virus membrane and that the C region, exposed to the interior face of the virion in a population of these mols., interacts with the nucleocapsid to which it is anchored, forming the core. Only the C region of the M protein is part of the core.

REFERENCE COUNT: 50 THERE ARE 50 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 11 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1999:336041 CAPLUS
 DOCUMENT NUMBER: 131:156705
 TITLE: Production, characterization, and uses of monoclonal antibodies against recombinant nucleoprotein of elk coronavirus
 AUTHOR(S): Daginakatte, Girish C.; Chard-Bergstrom, Cindy; Andrews, Gordon A.; Kapil, Sanjay
 CORPORATE SOURCE: Department of Diagnostic Medicine-Pathobiology, College of Veterinary Medicine, Manhattan, KS, 66506, USA
 SOURCE: Clinical and Diagnostic Laboratory Immunology (1999), 6(3), 341-344
 CODEN: CDIMEN; ISSN: 1071-412X
 PUBLISHER: American Society for Microbiology
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB This is the first report of the prodn. of monoclonal antibodies against elk coronavirus. The nucleoprotein gene of elk coronavirus was amplified by PCR and was cloned and expressed in a prokaryotic expression vector. Recombinant nucleocapsid protein was used to immunize mice for the prodn. of hybridomas. Twelve hybridomas that produced monoclonal antibodies against the nucleocapsid protein of elk coronavirus were selected by an indirect fluorescent-antibody test, an ELISA, and a Western blot assay. Ten of the monoclonal antibodies were of the IgG1 isotype, one was IgG2a, and one was IgM. All had kappa light chains. By immunohistochem. four monoclonal antibodies detected bovine coronavirus and elk coronavirus in formalin-fixed intestinal tissues. Anti-nucleoprotein monoclonal antibodies were better at ruminant coronavirus detection than the anti-spike protein monoclonal

antibodies. Because nucleoprotein is a more abundant antigen than spike protein in infected cells, this was not an unexpected finding.

REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 12 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1997:723088 CAPLUS
 DOCUMENT NUMBER: 128:58017
 ORIGINAL REFERENCE NO.: 128:11239a,11242a
 TITLE: Isolation and characterization of a coronavirus from elk calves with diarrhea
 AUTHOR(S): Majhdi, F.; Minocha, H. C.; Kapil, S.
 CORPORATE SOURCE: Department of Diagnostic Medicine/Pathobiology, College of Veterinary Medicine, Kansas State University, Manhattan, KS, 66506, USA
 SOURCE: Journal of Clinical Microbiology (1997), 35(11), 2937-2942
 CODEN: JCMIDW; ISSN: 0095-1137
 PUBLISHER: American Society for Microbiology
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB This is the first report of the isolation of a coronavirus from elk calves. Two fecal samples from elk calves with diarrhea were shown to be pos. for coronavirus-like particles by electron microscopy, and the particles were propagated in the human rectal tumor-18 cell line. After 24 h, syncytia were obsd., and cell culture supernatants from both samples showed hemagglutinating activity with mouse erythrocytes. Cells infected with both elk coronavirus (ECV) isolates reacted with Z3A5, a monoclonal antibody against the spike protein of bovine coronavirus (BCV), on an indirect fluorescent antibody test. The protein profiles of both ECV isolates were similar to that of BCV as detd. by sodium dodecyl sulfate-polyacrylamide gel electrophoresis anal. On Northern blot anal., the transcriptional pattern of ECV was typical of coronaviruses, with a nested set of transcripts with common 3' end sequences. Based on a published nucleoprotein gene sequence for BCV (Mebus isolate), we arbitrarily designed two primers for amplification by PCR. After cloning, the nucleoprotein was sequenced and a high degree of homol. (99%) between the nucleoprotein gene sequences of ECV and BCV was obsd. Thus, ECV is closely related genetically and antigenically to BCV and will be a new member of antigenic group 2 of the mammalian coronaviruses, which possess hemagglutinin-esterase protein.

L15 ANSWER 13 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1991:533524 CAPLUS
 DOCUMENT NUMBER: 115:133524
 ORIGINAL REFERENCE NO.: 115:22845a,22848a
 TITLE: Comparison of bovine coronavirus (BCV) antigens: monoclonal antibodies to the spike glycoprotein distinguish between vaccine and wild-type strains
 AUTHOR(S): Hussain, Khalid A.; Storz, Johannes; Kousoulas, Konstantin G.
 CORPORATE SOURCE: Sch. Vet. Med., Louisiana State Univ., Baton Rouge, LA, 70803, USA
 SOURCE: Virology (1991), 183(1), 442-5
 CODEN: VIRLAX; ISSN: 0042-6822
 DOCUMENT TYPE: Journal

LANGUAGE: English

AB Monoclonal antibodies (MAbs) against two major structural proteins of the cell-adapted Mebus strain of bovine coronavirus (BCV-L9) were produced and characterized. Seven MAbs reacted with the peplomeric glycoprotein, gp100/S, while three MAbs reacted with the nucleoprotein p53/N in Western blot anal. of BCV polypeptides. MAbs to gp100/S reacted with discontinuous epitopes of gp100/S in Westerns under mild but not under std. denaturing conditions. In contrast, MAbs to p53/N reacted in both types of Westerns, and those epitopes were thus continuous. MAbs to p53/N failed to neutralize BCV infectivity, while 4 MAbs to gp100/S neutralized BCV effectively. Cross reactivity of MAbs to gp100/S specified by five virulent wild-type strains and two high passage, cell-culture-adapted strains in mildly denaturing Westerns and neutralization assays indicated that two epitopes were conserved in all seven strains, while two epitopes of the avirulent strains were not detected in the wild-type strains. Non-neutralizing MAbs of gp100/S reacted with all seven strains in Westerns with the exception of one MAb that was specific for the highly cell-adapted strain BCV-L9.

L15 ANSWER 14 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1988:470029 CAPLUS
 DOCUMENT NUMBER: 109:70029
 ORIGINAL REFERENCE NO.: 109:11669a,11672a
 TITLE: Antigenic differentiation between transmissible gastroenteritis virus of swine and a related porcine respiratory coronavirus
 AUTHOR(S): Callebaut, P.; Correa, I.; Pensaert, M.; Jimenez, G.; Enjuanes, L.
 CORPORATE SOURCE: Fac. Vet. Med., State Univ. Gent, Ghent, B-9000, Belg.
 SOURCE: Journal of General Virology (1988), 69(7), 1725-30
 CODEN: JGVIAI; ISSN: 0022-1317
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB The antigenic relationship between isolated porcine respiratory coronavirus (TLM 83) and transmissible gastroenteritis (TGE) virus of swine was studied by neutralization, immunoblotting, and RIA, using TGE virus-specific monoclonal antibodies (MAbs) and polyclonal antibodies specific for both viruses. A complete two-way neutralization activity between the two viruses was found. Immunoblotting revealed cross-reactions between TLM 83 and TGE virus antigens at the level of the envelope protein (E1), the nucleoprotein (N), and the peplomer protein (E2). By virus neutralization assays and RIA with TGE virus-specific MAbs, the presence of similar epitopes in the E1 and N proteins and in the neutralization-mediating antigenic site of the E2 protein were demonstrated. E2 protein-specific MAbs, without neutralizing activity and reacting with antigenic sites B, C, and D (previously defined), failed to recognize TLM 83. These results indicated a close antigenic relationship and structural similarity between TLM 83 and TGE viruses and also suggested potential ways of differentiating between the two viruses.

L15 ANSWER 15 OF 15 CAPLUS COPYRIGHT 2009 ACS on STN



ACCESSION NUMBER: 1984:3195 CAPLUS
 DOCUMENT NUMBER: 100:3195
 ORIGINAL REFERENCE NO.: 100:551a,554a
 TITLE: Synthesis and subcellular localization of the murine coronavirus nucleocapsid protein

AUTHOR(S): Stohlman, Stephen A.; Fleming, John O.; Patton, Chris
D.; Lai, Michael M. C.
CORPORATE SOURCE: Sch. Med., Univ. South. California, Los Angeles, CA,
90033, USA
SOURCE: Virology (1983), 130(2), 527-32
CODEN: VIRLAX; ISSN: 0042-6822
DOCUMENT TYPE: Journal
LANGUAGE: English

AB The synthesis and processing of the nucleocapsid protein (pp60) of the JHM strain of murine coronaviruses were examd. Pulse-chase expts. showed that pp60 was synthesized initially as a protein of mol. wt. ~57,000 (p57). Immunopptn. using mouse anti-JHMV antiserum indicated that p57 was virus specific. Immunopptn. with monoclonal antibodies specific for pp60 showed that p57 was antigenically related to pp60 and was not phosphorylated, whereas the intracellular protein that comigrated with the virion nucleocapsid protein, pp60, was phosphorylated. The p57 was found exclusively in the cytosol whereas the majority of pp60 was assocd. with the membrane fraction but pp60 was not an integral membrane protein.

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